



1 16.(previously presented) The composition of claim 10, wherein each of the monomers  
2 comprises a deoxynucleotide triphosphate (dNTP) and the monomer tag is covalently bonded to the  
3  $\beta$  or  $\gamma$  phosphate group of each dNTP.

1 17.(previously presented) The composition of claim 10, wherein the tags comprise fluorescent  
2 tags and the fluorescence property comprises an intensity and/or frequency of emitted fluorescent  
3 light.

1 18.(previously presented) The composition of claim 17, wherein the fluorescence property is  
2 fluorescence resonance energy transfer (FRET) where either the monomer tag or the polymerase tag  
3 comprises a donor and the other tag comprises an acceptor and where FRET occurs when the two  
4 tags are in close proximity.

5 19.(previously presented) The composition of claim 14, wherein the polymerase comprises *Taq*  
6 DNA polymerase I having a tag attached to an amino acid at a specific amino acid position of the  
7 *Taq* DNA polymerase I, where the amino acid position is selected from the group consisting of 513-  
8 518, 643, 647, 649 and 653-661 of SEQ. ID No. 11, where the tag comprises a fluorescent molecule.

20.(canceled)

21.(canceled)

22.(canceled)

22.(canceled)

23.(canceled)

24.(canceled)

1 25.(withdrawn) A single molecule sequencing apparatus comprising a substrate having a first  
2 chamber in which at least one tagged polymerase is confined therein and a second chamber including  
3 tagged dNTPs and a channel interconnecting the chambers, where a detectable property of at least  
4 one tag undergoes a detectable change during a monomer incorporation cycle.

1 26.(withdrawn) The apparatus of claims 24, further comprising a plurality of monomer  
2 chambers, one for each tagged dNTP.

1 27.(withdrawn) A mutant Taq polymerase comprising native Taq polymerase with a cysteine  
2 residue replacement at a site selected from the group consisting of 513-518, 643, 647, 649 and 653-  
3 661 and mixtures or combinations thereof.

1 28.(withdrawn) The polymerase of claim 27, wherein the cysteine residue includes a tag  
2 covalently bonded thereto through the SH group.

1 29.(withdrawn) A system for retrieving stored information comprising:  
2 a unknown nucleotide sequence representing a data stream;  
3 a single-molecule sequencer including a polymerase having a tag associated therewith and  
4 monomers for the polymerase, each monomer having a tag associated therewith;  
5 an excitation source adapted to excite the at least one of the tags; and  
6 a detector adapted to detect a response from at least one of the tag,  
7 where the response changes during polymerization of a complementary sequence and the  
8 changes in response represent a content of the data stream.

1 30.(withdrawn) A system for determining sequence information from a single molecule  
2 comprising:  
3 a unknown nucleotide sequence;  
4 a single-molecule sequencer comprising a polymerase having a tag associated therewith and  
5 monomers for the polymerase, each monomer having a tag associated therewith;  
6 a excitation source adapted to excite at least one of the tags; and  
7 a detector adapted to detect a response from at least one of the tags,  
8 where the response changes during polymerization of a complementary sequence and the  
9 changes in the response represent the identity of each nucleotide in the unknown sequence.

1 31.(withdrawn) A method for sequencing a molecular sequence comprising:  
2 supplying an unknown sequence of nucleotides or nucleotide analogs to a single-molecule

3 sequencer comprising a polymerase having a fluorescent donor covalently attached thereto and  
4 monomers for the polymerase, each monomer having a unique fluorescent acceptor covalently  
5 bonded to the beta or gamma phosphate thereof;

6 exciting the fluorescent donor with a light from an excitation light source;

7 detecting emitted fluorescent light from the acceptor during a monomer incorporation cycle  
8 via a fluorescent light detector, where an intensity and/or frequency of the emitted light for the  
9 incorporating acceptors changes during each monomer incorporation cycle; and

10 converting the changes into an identity of each nucleotide or nucleotide analog in the  
11 unknown sequence.

1 32.(withdrawn) A method of sequencing an individual nucleic acid molecule or numerous  
2 individual molecules in parallel including the steps of:

3 immobilizing a member of the replication complex comprising a polymerase including a tag  
4 attached thereto, a primer or a template sufficiently spaced apart to allow resolution detection of each  
5 complex on a solid support;

6 incubating the replication complex with cooperatively-tagged nucleotides, each nucleotide  
7 including a unique tag at its gamma-phosphate, where each nucleotide can be individually detected;

8 detecting each nucleotide incorporated by the polymerase as the polymerase transitions  
9 between its open and closed form, which causes a change in a detectable property of at least one of  
10 the tags or as the pyrophosphate group is released by the polymerase; and

11 relating the changes in the detectable property to the sequence of nucleotides in an unknown  
12 nucleic acid sequence.

1 33.(withdrawn) A  $\gamma$ -phosphate modified nucleoside comprising  $\gamma$ -phosphate modified dATP,  
2 dCTP, dGTP and dTTP.

1 34.(withdrawn) A primer sequence or portion thereof selected from the group consisting of  
2 Sequence 1 through 29.

35.(canceled)

36.(canceled)



1 54.(previously presented) The composition of claim 50, wherein the tags comprise fluorescent  
2 tags and the fluorescence property comprises an intensity and/or frequency of emitted fluorescent  
3 light.

1 55.(previously presented) The composition of claim 54, wherein the fluorescence property is  
2 fluorescence resonance energy transfer (FRET) where either the monomer tag or the polymerase tag  
3 comprises a donor and the other tag comprises an acceptor and where FRET occurs when the two  
4 tags are in close proximity.

5 56.(previously presented) The composition of claim 52, wherein the polymerase comprises *Taq*  
6 DNA polymerase I having a tag attached to an amino acid at a specific amino acid position of the  
7 *Taq* DNA polymerase I, where the amino acid position is selected from the group consisting of 513-  
8 518, 643, 647, 649 and 653-661 of SEQ. ID No. 11, where the tag comprises a fluorescent molecule.